DATE: 10/30/2001

TIME: 12:01:27

OIPE

```
PATENT APPLICATION: US/09/903,023
                     Input Set : A:\Rih32dll.app
                     Output Set: N:\CRF3\10302001\I903023.raw
      3 <110> APPLICANT: Wands, Jack R.
             de la Monte, Suzanne M.
     5
             Ince, Nedim
             Carlson, Rolf I.
     8 <120> TITLE OF INVENTION: DIAGNOSIS AND TREATMENT OF MALIGNANT NEOPLASMS
     10 <130> FILE REFERENCE: 21486-032 DIV1
     12 <140> CURRENT APPLICATION NUMBER: 09/903,023
C--> 13 <141> CURRENT FILING DATE: 2001-10-11
     15 <150> PRIOR APPLICATION NUMBER: 09/436,184
     16 <151> PRIOR FILING DATE: 1999-11-08
                                                                 ENTERED
     18 <160> NUMBER OF SEQ ID NOS: 9
     20 <170> SOFTWARE: PatentIn Ver. 2.1
     22 <210> SEQ ID NO: 1
     23 <211> LENGTH: 36
     24 <212> TYPE: PRT
     25 <213> ORGANISM: Artificial Sequence
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     28 <223> OTHER INFORMATION: Description of Artificial Sequence: Consensus
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     33 <222> LOCATION: (2)..(8)
     34 <223> OTHER INFORMATION: Wherein Xaa is any amino acid
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     38 <222> LOCATION: (10)..(13)
     39 <223> OTHER INFORMATION: Wherein Xaa is any amino acid.
     41 <220> FEATURE:
     42 <221> NAME/KEY: VARIANT
     43 <222> LOCATION: (15)..(24)
     44 <223> OTHER INFORMATION: Wherein Xaa is any amino acid. \sim
     46 <220> FEATURE:
     47 <221> NAME/KEY: VARIANT
     48 <222> LOCATION: (26)
     49 <223> OTHER INFORMATION: Wherein Xaa is any amino acid.
     51 <220> FEATURE:
     52 <221> NAME/KEY: VARIANT
     53 <222> LOCATION: (28)..(35)
     54 <223> OTHER INFORMATION: Wherein Xaa is any amino acid.
     56 <400> SEQUENCE:
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W--> 60 Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Cys (Xaa Cys (Xaa Xaa Xaa Xaa Xaa
                                                             30
    61
W--> 63 Xaa Xaa Xaa Cys
             ---35
     64
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RAW SEQUENCE LISTING

67 <210> SEQ ID NO: 2

RAW SEQUENCE LISTING DATE: 10/30/2001 PATENT APPLICATION: US/09/903,023 TIME: 12:01:27

Input Set : A:\Rih32dll.app

Output Set: N:\CRF3\10302001\I903023.raw

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70 <213> ORGANISM: Homo sapiens
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76 Gly Ser Gly Ser Gly Ser Thr Ser Ala Gly Ser Ser Pro Gly Ala
79 Arg Arg Glu Thr Lys His Gly Gly His Lys Asn Gly Arg Lys Gly Gly
                               40
82 Leu Ser Gly Thr Ser Phe Phe Thr Trp Phe Met Val Ile Ala Leu Leu
85 Gly Val Trp Thr Ser Val Ala Val Val Trp Phe Asp Leu Val Asp Tyr
                       70
88 Glu Glu Val Leu Gly Lys Leu Gly Ile Tyr Asp Ala Asp Gly Asp Gly
                   85
91 Asp Phe Asp Val Asp Asp Ala Lys Val Leu Leu Gly Leu Lys Glu Arg
              100
                                  105
94 Ser Thr Ser Glu Pro Ala Val Pro Pro Glu Glu Ala Glu Pro His Thr
                              120
97 Glu Pro Glu Glu Gln Val Pro Val Glu Ala Glu Pro Gln Asn Ile Glu
                          135
100 Asp Glu Ala Lys Glu Gln Ile Gln Ser Leu Leu His Glu Met Val His
                       150
                                           155
103 Ala Glu His Val Glu Gly Glu Asp Leu Gln Glu Asp Gly Pro Thr
                                       170
                   165
106 Gly Glu Pro Gln Gln Glu Asp Asp Glu Phe Leu Met Ala Thr Asp Val
                                   185
               180
109 Asp Asp Arg Phe Glu Thr Leu Glu Pro Glu Val Ser His Glu Glu Thr
110 , 195
                               200
112 Glu His Ser Tyr His Val Glu Glu Thr Val Ser Gln Asp Cys Asn Gln
                           215
                                               220
115 Asp Met Glu Glu Met Met Ser Glu Gln Glu Asn Pro Asp Ser Ser Glu
                       230
                                           235
118 Pro Val Val Glu Asp Glu Arg Leu His His Asp Thr Asp Asp Val Thr
                   245
121 Tyr Gln Val Tyr Glu Glu Gln Ala Val Tyr Glu Pro Leu Glu Asn Glu
               260
                                   265
124 Gly Ile Glu Ile Thr Glu Val Thr Ala Pro Pro Glu Asp Asn Pro Val
                               280
127 Glu Asp Ser Gln Val Ile Val Glu Glu Val Ser Ile Phe Pro Val Glu
                                               300
128 290
                           295
130 Glu Gln Gln Glu Val Pro Pro Glu Thr Asn Arg Lys Thr Asp Asp Pro
                                           315
                       310
133 Glu Gln Lys Ala Lys Val Lys Lys Lys Pro Lys Leu Leu Asn Lys
                   325
                                       330
136 Phe Asp Lys Thr Ile Lys Ala Glu Leu Asp Ala Ala Glu Lys Leu Arg
                                   345
139 Lys Arg Gly Lys Ile Glu Glu Ala Val Asn Ala Phe Lys Glu Leu Val
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RAW SEQUENCE LISTING DATE: 10/30/2001 PATENT APPLICATION: US/09/903,023 TIME: 12:01:27

Input Set : A:\Rih32dll.app

Output Set: N:\CRF3\10302001\1903023.raw

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143	•	370	•				375			_	-	380	•			•
145	Glu	Asp	Asp	Leu	Ala	Glu	Lys	Arg	Arg	Ser	Asn	Glu	Val	Leu	Arg	Gly
	385	_	_			390	_	_	_		395				_	400
148	Ala	Ile	Glu	Thr	Tyr	Gln	Glu	Val	Ala	Ser	Leu	Pro	Asp	Val	Pro	Ala
149					405					410			_		415	
151	Asp	Leu	Leu	Lys	Leu	Ser	Leu	Lys	Arg	Arg	Ser	Asp	Arg	Gln	Gln	Phe
152	-			420				-	425	-		_	_	430		
154	Leu	Gly	His	Met	Arg	Gly	Ser	Leu	Leu	Thr	Leu	Gln	Arg	Leu	Val	Gln
155		_	435		_	_		440					445			
157	Leu	Phe	Pro	Asn	Asp	Thr	Ser	Leu	Lys	Asn	Asp	Leu	Gly	Val	Gly	Tyr
158		450					455					460				
160	Leu	Leu	Ile	Gly	Asp	Asn	Asp	Asn	Ala	Lys	Lys	Val	Tyr	Glu	Glu	Val
161	465					470					475					480
163	Leu	Ser	Val	Thr	Pro	Asn	Asp	Gly	Phe	Ala	Lys	Val	His	Tyr	Gly	Phe
164					485					490					495	
166	Ile	Leu	Lys	Ala	Gln	Asn	Lys	Ile	Ala	Glu	Ser	Ile	Pro	Tyr	Leu	Lys
167				500					505					510		
169	Glu	Gly	Ile	Glu	Ser	Gly	Asp	Pro	Gly	Thr	Asp	Asp	Gly	Arg	Phe	Tyr
170			515					520					525			
172	Phe	His	Leu	Gly	Asp	Ala	Met	Gln	Arg	Val	Gly	Asn	Lys	Glu	Ala	${ t Tyr}$
173		530					535					540				
175	Lys	\mathtt{Trp}	Tyr	Glu	Leu	Gly	His	Lys	Arg	Gly	His	Phe	Ala	Ser	Val	\mathtt{Trp}
	545					550					555					560
178	Gln	Arg	Ser	Leu	Tyr	Asn	Val	Asn	Gly	Leu	Lys	Ala	Gln	Pro		\mathtt{Trp}
179					565					570					575	
	Thr	Pro	Lys		Thr	Gly	Tyr	Thr		Leu	Val	Lys	Ser		Glu	Arg
182				580				•	585	_		•		590	_	
	Asn	Trp	_	Leu	Ile	Arg	Asp		Gly	Leu	Ala	Val		Asp	Lys	Ala
185	_	~-	595	1	_	_		600	~ 1		-0	_	605	_	a 3	
	Lys	_	Leu	Phe	Leu	Pro		Asp	GLu	Asn	Leu		GLu	Lys	GIY	Asp
188	_	610	-1	-1	-1	_	615	61	01	a 1		620	•	a 3		
	Trp	Ser	GIn	Pne	Thr		Trp	GIn	GIn	GLY	_	Arg	Asn	GIU	Asn	
	625	•	a 1	. 1 -	D	630	m 1	G	m 1	T	635	01	T	Dha	D===	640
	Cys	гаг	GIY	Ата	645	гуѕ	THE	Cys	THE	650	Leu	GIU	гаг	Pile	655	GIU
194	m 1	m1	a1	0		7	C1	01 =	T1.		m	Com	T1.	Wot		Dwo
	Thr	Thr	GIY		Arg	Arg	GTĀ	GIII		гаг	TYL	ser	TTE	670	нтѕ	PIO
197	Gly	mh m	114.0	660	mzz	Dro	uic	mb~	665	Dro	mh~	Nan	Cvc		Lou	λνα
	СТА	THE	675	vaı	ттр	PIO	HIS	680	GIY	PIO	TIII	ASII	685	AIG	ьeu	AIG
200	Mat	111.0		c1	T 011	370]	т1 о		T	C1	C1	Crra		т1 о	λνα	Cvc
	Met		ьeu	сту	ьeu	val	695	PIO	nys	GIU	GIĀ	700	пур	TTG	мту	Cys
203	Ala	690	C1	πh∽	λ ~~	mb~		<i>c</i> 1	G1 11	C117	Lvc		Leu	Tle	Dhe	Δen
	705	หอแ	GIU	TIIT	ary	710	тъ	GIU	GLU	ату	715	val	пeп	TT6	r me	720
	Asp	Ser	Dhe	6111	Wie		Va 1	ጥፖጥ	Gln	Δen		Ser	Ser	Phe	Arσ	
209	vəħ	JEI	FIIG	GIU	725	GIU	, a i	115	0111	730	ru	JCI	JUL	1	735	
	Ile	Dho	٦١۵	Val		Va 1	ጥተካ	ніс	Pro		Len	Thr	Pro	Gln		Ara
212	***	1 110	110	740	5	, 41			745	O_Lu	u			750		5
212				, 40					, 13							

RAW SEQUENCE LISTING DATE: 10/30/2001 PATENT APPLICATION: US/09/903,023 TIME: 12:01:27

Input Set : A:\Rih32dll.app

Output Set: N:\CRF3\10302001\I903023.raw

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221 <213> ORGANISM: Homo sapiens
223 <400> SEQUENCE: 3
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226 agcatggagg acacaagaat gggaggaaag gcggactctc gggaacttca ttcttcacgt 180
227 ggtttatggt gattgcattg ctgggcgtct ggacatctgt agctgtcgtt tggtttgatc 240
228 ttgttgacta tgaggaagtt ctaggaaaac taggaatcta tgatgctgat ggtgatggag 300
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230 caqcaqtccc gccagaagag gctgagccac acactgagcc cgaggagcag gttcctgtgg 420
231 aggcagaacc ccagaatatc gaagatgaag caaaagaaca aattcagtcc cttctccatg 480
232 aaatggtaca cgcagaacat gttgagggag aagacttgca acaagaagat ggacccacag 540
233 gagaaccaca acaagaggat gatgagtttc ttatggcgac tgatgtagat gatagatttg 600
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235 cagtttcaca agactgtaat caggatatgg aagagatgat gtctgagcag gaaaatccag 720
236 attccagtga accagtagta gaagatgaaa gattgcacca tgatacagat gatgtaacat 780
237 accaagtcta tgaggaacaa gcagtatatg aacctctaga aaatgaaggg atagaaatca 840
238 cagaagtaac tgctcccct gaggataatc ctgtagaaga ttcacaggta attgtagaag 900
239 aagtaagcat ttttcctgtg gaagaacagc aggaagtacc accagaaaca aatagaaaaa 960
240 cagatgatee agaacaaaaa gcaaaagtta agaaaaagaa geetaaaett ttaaataaat 1020
241 ttgataagac tattaaagct gaacttgatg ctgcagaaaa actccgtaaa aggggaaaaa 1080
242 ttgaggaagc agtgaatgca tttaaagaac tagtacgcaa ataccctcag agtccacgag 1140
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259 aggaaggetg caagattega tgtgecaaeg agaceaggae etgggaggaa ggeaaggtge 2160
260 tcatctttga tgactccttt gagcacgagg tatggcagga tgcctcatct ttccggctga 2220
261 tattcatcgt ggatgtgtgg catccggaac tgacaccaca gcagagacgc agccttccag 2280
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265 <210> SEQ ID NO: 4
266 <211> LENGTH: 31
267 <212> TYPE: PRT
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DATE: 10/30/2001

TIME: 12:01:27

Input Set : A:\Rih32dll.app Output Set: N:\CRF3\10302001\1903023.raw 268 <213> ORGANISM: Artificial Sequence 270 <220> FEATURE: 271 <223> OTHER INFORMATION: Description of Artificial Sequence: EGF-like 272 cysteine-rich repeat 274 <220> FEATURE: 275 <221> NAME/KEY: VARIANT 276 <222> LOCATION: (3)..(5) 277 <223> OTHER INFORMATION: Wherein any Xaa may be any amino acid 279 <220> FEATURE: 280 <221> NAME/KEY: VARIANT 281 <222> LOCATION: (6)..(7) 282 <223> OTHER INFORMATION: Wherein Xaa is any amino acid. 284 <220> FEATURE: 285 <221> NAME/KEY: VARIANT 286 <222> LOCATION: (10) 287 <223> OTHER INFORMATION: Wherein Xaa is any amino acid. 289 <220> FEATURE: 290 <221> NAME/KEY: VARIANT 291 <222> LOCATION: (14) 292 <223> OTHER INFORMATION: Wherein Xaa is any amino acid. 294 <220> FEATURE: 295 <221> NAME/KEY: VARIANT 296 <222> LOCATION: (17)..(18) 298 <220> FEATURE: 299 <221> NAME/KEY: VARIANT 300 <222> LOCATION: (25)..(26) 301 <223> OTHER INFORMATION: Wherein Xaa is any amino acid. 303 <220> FEATURE: 304 <221> NAME/KEY: VARIANT 305 <222> LOCATION: (29) 306 <223> OTHER INFORMATION: Wherein Xaa is any amino acid. 308 <400> SEQUENCE: 4 W--> 309 Cys Asp Xaa Xaa Xaa Cys Xaa Xaa Lys Xaa Gly Asn Gly Xaa Cys Asp 310/1 ______5 W--> 312 Xaa Xaa Cys Asn Asn Ala Ala Cys Xaa Xaa Asp Gly Xaa Asp Cys 313 316 <210> SEQ ID NO: 5 317 <211> LENGTH: 1242 318 <212> TYPE: PRT 319 <213> ORGANISM: Homo sapiens 321 <400> SEQUENCE: 5 322 Met Ala Ser Pro Pro Glu Ser Asp Gly Phe Ser Asp Val Arg Lys Val 323 10 325 Gly Tyr Leu Arg Lys Pro Lys Ser Met His Lys Arg Phe Phe Val Leu 328 Arg Ala Ala Ser Glu Ala Gly Gly Pro Ala Arg Leu Glu Tyr Tyr Glu 40 331 Asn Glu Lys Lys Trp Arg His Lys Ser Ser Ala Pro Lys Arg Ser Ile 332

RAW SEQUENCE LISTING

PATENT APPLICATION: US/09/903,023

VERIFICATION SUMMARY

PATENT APPLICATION: US/09/903,023

DATE: 10/30/2001

TIME: 12:01:28

Input Set : A:\Rih32dll.app

Output Set: N:\CRF3\10302001\I903023.raw

L:13 M:271 C: Current Filing Date differs, Replaced Current Filing Date

L:57 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:1
L:60 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:1
L:63 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:1
L:309 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:4
L:312 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:4